Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1. (previously presented) A method of large scale virus production, which comprises:

a) inoculating a cell growth medium with a population of mammalian host cells, wherein the cell growth medium contains an effective amount of a shear-protective compound;

b) culturing the mammalian host cells in the cell growth medium;

c) infecting the mammalian host cells in the cell growth medium with an aliquot of a virus seed stock, wherein the virus seed is essentially free of any cell-lysing component;

d) culturing the virus infected mammalian host cells of step c) under gas sparging;

e) harvesting intracellular and/or extracellular virus from the mammalian host cells and cell growth medium; and,

f) purifying the harvested virus of step e);

wherein the shear-protective compound is a block copolymer surfactant.

2. (canceled)

3. (currently amended) The method of claim 1 wherein the shear-protective compound is ~~PLURONIC®F-68~~ a polyoxyethylene-polyoxypropylene block copolymer having an average molecular weight of 8400 Da.

4. (currently amended) The method of claim 3 wherein ~~PLURONIC®F-68~~ the polyoxyethylene-polyoxypropylene block copolymer is present at a concentration from about 0.3 g/L to about 10 g/L.

5. (currently amended) The method of claim 4 wherein ~~PLURONIC®F-68~~ the polyoxyethylene-polyoxypropylene block copolymer is present at a concentration from about 1 g/L to about 2 g/L.

6. (previously presented) The method of claim 1 wherein gas sparging is provided at a rate up to about 0.1 VVM.

7. (previously presented) The method of claim 6 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

8-9. (canceled)

10. (previously presented) The method of claim 3 wherein gas sparging is provided at a rate up to about 0.1 VVM.

11. (previously presented) The method of claim 10 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

12. (previously presented) The method of claim 4 wherein gas sparging is provided at a rate up to about 0.1 VVM.

13. (previously presented) The method of claim 12 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

14. (previously presented) The method of claim 5 wherein gas sparging is provided at a rate up to about 0.1 VVM.

15. (previously presented) The method of claim 14 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

16. (previously presented) A method of large scale adenovirus production, which comprises:

a) inoculating a cell growth medium with a population of mammalian host cells, wherein the cell growth medium contains an effective amount of a shear-protective compound;

b) culturing the mammalian host cells in the cell growth medium;

c) infecting the mammalian host cells in the cell growth medium with an aliquot of an adenovirus seed stock, wherein the adenovirus seed is essentially free of any cell-lysing component;

d) culturing the adenovirus infected mammalian host cells of step c) under gas sparging;

e) harvesting intracellular and/or extracellular adenovirus from the mammalian host cells and cell growth medium; and,

f) purifying the harvested adenovirus of step e);

wherein the shear-protective compound is a block copolymer surfactant.

17. (canceled)

18. (currently amended) The method of claim 16 wherein the shear-protective compound is ~~PLURONIC®F-68~~ a polyoxyethylene-polyoxypropylene block copolymer having an average molecular weight of 8400 Da.

19. (currently amended) The method of claim 18 wherein ~~PLURONIC®F-68~~ the polyoxyethylene-polyoxypropylene block copolymer is present at a concentration from about 0.3 g/L ~~and~~ to about 10 g/L.

20. (currently amended) The method of claim 19 wherein ~~PLURONIC®F-68~~ the polyoxyethylene-polyoxypropylene block copolymer is present at a concentration from about 1 g/L to about 2 g/L.

21. (previously presented) The method of claim 16 wherein gas sparging is provided at a rate up to about 0.1 VVM.

22. (previously presented) The method of claim 21 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

23-24. (canceled)

25. (previously presented) The method of claim 18 wherein gas sparging is provided at a rate up to about 0.1 VVM.

26. (previously presented) The method of claim 25 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

27. (previously presented) The method of claim 19 wherein gas sparging is provided at a rate up to about 0.1 VVM.

28. (previously presented) The method of claim 27 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

29. (previously presented) The method of claim 20 wherein gas sparging is provided at a rate up to about 0.1 VVM.

30. (previously presented) The method of claim 29 wherein gas sparging is provided at a rate up to about 0.001 to 0.05 VVM.

31-46. (canceled)

47. (previously presented) The method of claim 16 wherein the mammalian host cells are PER.C6™ E1-complementing cells.

48. (new) The method of claim 1 wherein the cell-lysing component is present at a concentration less than 0.00025%.

49. (new) The method of claim 16 wherein the cell-lysing component is present at a concentration less than 0.00025%.